

PI Newton DL, Rybak SM;
XX
XX WPI: 1999-610847/52.
DR N-PSDB: AA208125.
XX
PT New recombinant ribonucleases, used for killing target cells, e.g. for
PT treating cancers, viral infections or autoimmune diseases
XX
PS Claim 34; Page 56; 71pp: English.
XX
CC The present sequence is a recombinant Rana pipiens ribonuclease (RapLRI)
CC protein with Met23Leu. Carboxy terminal end of recombinant RapLRI has a
CC covalently bound ligand binding moiety, which can be a LL2 antibody
CC directed against CD22 on cancerous B cells or human chorionic
CC gonadotropin (hCG) effective against Kaposi's sarcoma cells. Recombinant
CC ribonucleases can be expressed in bacteria without an N-terminal
CC methionine due to the presence of a signal peptide that is cleaved by
CC bacteria. The soluble expression of ribonuclease allows the proteins to
CC be fused in-frame with ligand binding moieties to form cytotoxic fusion
CC proteins. They can be used for treatment of cancer and autoimmune
CC diseases.
XX
SQ Sequence 104 AA;
Query Match 98.6%; Score 571; DB 20; Length 104;
Best Local Similarity 99.0%; Pred. No. 2.4e-62;
Matches 103; Conservative 0; Mismatches 1; Indels 0; Gaps 0;
OY 1 QDWLTFQKHLNTRDVCNNILSTNLFHCKDKMTFTYSRPEPYKAICKGIASKNVLT 60
Db 1 QDWLTFQKHLNTRDVCNNILSTNLFHCKDKMTFTYSRPEPYKAICKGIASKNVLT 60
OY 61 FEFLSDCQNTSRPCRYKRLKSTNTPFCVTCENQAPVHFVGHC 104
Db 61 SEFLSDCQNTSRPCRYKRLKSTNTPFCVTCENQAPVHFVGHC 104
RESULT 2
AAV28869 ID AAV28869 standard; Protein: 105 AA.
XX
AC AAV28869;
XX
DT 25-JAN-2000 (first entry)
XX
DE Recombinant Met(-1) RapLRI Met23Leu-(His)6 protein.
XX
KW Recombinant Met(-1) Rana pipiens ribonuclease Met23Leu-(His)6; RapLRI;
KW CD22; covalently bound; LL2 antibody; ligand binding moiety; RNase;
KW cancerous B cell; Kaposi's sarcoma; human chorionic gonadotropin; hCG;
KW signal peptide; recombinant ribonuclease; cytotoxic fusion protein;
KW cancer; frog; autoimmune disease.
XX
OS Rana pipiens.
OS Synthetic.
XX
FH Key Location/Qualifiers
FH MISC-difference 1 /note= "(His)6 histidine tag attached to N-terminal Met"
FT MISC-difference 1 /note= "Met not found in wild type RapLRI"
FT MISC-difference 24 /note= "Wild type Met replaced with Leu"
FT
XX
XX WO950398-A2.
XX
XX 07-OCT-1999.
XX
XX 26-MAR-1999; 99WO-US06641.
XX
XX 27-MAR-1998; 98US-0079751.
XX
XX (USSH) US DEPT HEALTH & HUMAN SERVICES.
XX

XX
XX Newton DL, Rybak SM;
XX
XX WPI: 1999-610847/52.
DR N-PSDB: AA208127.
XX
PT New recombinant ribonucleases, used for killing target cells, e.g. for
PT treating cancers, viral infections or autoimmune diseases
XX
PS Claim 4; Page 59; 71pp: English.
XX
CC The present sequence is a recombinant Rana pipiens ribonuclease protein
CC (RapLRI) with Met at position 1 attached to (His)6 tag and Met24Leu.
CC Carboxy terminal end of recombinant RapLRI has a covalently bound ligand
CC binding moiety, which can be a LL2 antibody directed against CD22 on
CC cancerous B cells or human chorionic gonadotropin (hCG) effective
CC against Kaposi's sarcoma cells. Recombinant ribonucleases can be
CC expressed in bacteria without an N-terminal methionine due to the
CC presence of a signal peptide that is cleaved by bacteria. The soluble
CC expression of ribonuclease allows the proteins to be fused in-frame with
CC ligand binding moieties to form cytotoxic fusion proteins. They can be
CC used for treatment of cancer and autoimmune diseases.
XX
SQ Sequence 105 AA;
Query Match 98.6%; Score 571; DB 20; Length 105;
Best Local Similarity 99.0%; Pred. No. 2.4e-62;
Matches 103; Conservative 0; Mismatches 1; Indels 0; Gaps 0;
OY 1 QDWLTFQKHLNTRDVCNNILSTNLFHCKDKMTFTYSRPEPYKAICKGIASKNVLT 60
Db 2 QDWLTFQKHLNTRDVCNNILSTNLFHCKDKMTFTYSRPEPYKAICKGIASKNVLT 61
OY 61 FEFLSDCQNTSRPCRYKRLKSTNTPFCVTCENQAPVHFVGHC 104
Db 62 SEFLSDCQNTSRPCRYKRLKSTNTPFCVTCENQAPVHFVGHC 105
RESULT 3
AAV28865 ID AAV28865 standard; Protein: 104 AA.
XX
AC AAV28865;
XX
DT 25-JAN-2000 (first entry)
XX
DE Rana pipiens liver ribonuclease (RapLRI).
XX
KW Rana pipiens liver ribonuclease; RapLRI; covalently bound; LL2 antibody;
KW ligand binding moiety; CD22; cancerous B cell; Kaposi's Sarcoma; frog;
KW human chorionic gonadotropin; hCG; recombinant ribonuclease; RNase;
KW signal peptide; cytotoxic fusion protein; cancer; autoimmune disease.
XX
OS Rana pipiens.
OS Synthetic.
XX
FH WO950398-A2.
FH 07-OCT-1999.
PD
XX
XX 26-MAR-1999; 99WO-US06641.
XX
XX 27-MAR-1998; 98US-0079751.
XX
XX (USSH) US DEPT HEALTH & HUMAN SERVICES.
XX
XX Newton DL, Rybak SM;
XX
XX WPI: 1999-610847/52.
DR N-PSDB: AA208124.
XX
XX
PT New recombinant ribonucleases, used for killing target cells, e.g. for
PT treating cancers, viral infections or autoimmune diseases
XX

PS Claim 1; Page 55; 71pp; English.

XX The present sequence is Rana pipiens liver ribonuclease (RapLr1)

CC protein. Carboxy terminal end of RapLr1 has a covalently bound

CC ligand binding moiety, which can be a LL2 antibody directed against

CC CD22 on cancerous B cells or human chorionic gonadotropin (hCG)

CC effective against Kaposi's Sarcoma cells. Recombinant ribonucleases can

CC be expressed in bacteria without an N-terminal methionine due to the

CC presence of a signal peptide that is cleaved by bacteria. The soluble

CC expression of ribonuclease allows the proteins to be fused in-frame with

CC ligand binding moieties to form cytotoxic fusion proteins. They can be

CC used for treatment of cancer and autoimmune diseases.

XX

SQ Sequence 104 AA:

Query Match 98.3%; Score 569; DB 20; Length 104;

Best Local Similarity 98.1%; Pred. No. 4.2e-62;

Matches 102; Conservative 1; Mismatches 1; Indels 0; Gaps 0;

OY 1 QDWLTFQKKHLTNRVDVDCNNILSTNLFHCKDKNTFYSPPEPKAICGIIASKNVLTT 60

DB 1 QDWLTFQKKHLTNRVDVDCNNIMSTNLFHCKDKNTFYSPPEPKAICGIIASKNVLTT 60

OY 61 FEFLSDCNVTSRPCKYKLLKSTNFCVTCENQAPVHFVGHC 104

DB 61 SEFLSDCNVTSRPCKYKLLKSTNFCVTCENQAPVHFVGHC 104

RESULT 4

AAV28867

ID AAV28867 standard; Protein; 105 AA.

AC AAV28867;

XX 25-JAN-2000 (first entry)

DT

DE Recombinant Met(-1) RapLr1.

XX

KM Recombinant Met(-1) Rana pipiens ribonuclease; RapLr1; CD22; RNase;

KM covalently bound; LL2 antibody; ligand binding moiety; cancerous B cell;

KM Kaposi's Sarcoma; human chorionic gonadotropin; hCG; signal peptide;

KM recombinant ribonuclease; cytotoxic fusion protein; cancer; frog;

KW autoimmune disease.

XX

OS Rana pipiens.

OS Synthetic.

OS

FT Key Location/Qualifiers

FT MISC-difference 1 /note="Met not found in wild type RapLr1"

XX

PN W09950398-A2.

XX

PD 07-OCT-1999.

XX

XX 26-MAR-1999; 99WO-US06641.

PF

XX 27-MAR-1998; 98US-0079751.

PR

XX (USSH) US DEPT HEALTH & HUMAN SERVICES.

PA

PI Newton DL, Rybak SM;

XX

DR WPI: 1999-610847/52.

DR N-PSDB; AAZ08126.

XX

XX New recombinant ribonucleases, used for killing target cells, e.g. for

PT treating cancers, viral infections or autoimmune diseases

XX

PS Claim 34; Page 57; 71pp; English.

XX

CC The present sequence is a recombinant Rana pipiens ribonuclease (RapLr1)

CC protein with Met at position 1. Carboxy terminal end of recombinant

CC RapLr1 has a covalently bound ligand binding moiety, which can be a LL2

CC antibody directed against CD22 on cancerous B cells or human chorionic

CC gonadotropin (hCG) effective against Kaposi's sarcoma cells. Recombinant

CC ribonucleases can be expressed in bacteria without an N-terminal

CC methionine due to the presence of a signal peptide that is cleaved by

CC bacteria. The soluble expression of ribonuclease allows the proteins to

CC be fused in-frame with ligand binding moieties to form cytotoxic fusion

CC proteins. They can be used for treatment of cancer and autoimmune

CC diseases.

XX

SQ Sequence 105 AA:

Query Match 98.3%; Score 569; DB 20; Length 105;

Best Local Similarity 98.1%; Pred. No. 4.3e-62;

Matches 102; Conservative 1; Mismatches 1; Indels 0; Gaps 0;

OY 1 QDWLTFQKKHLTNRVDVDCNNILSTNLFHCKDKNTFYSPPEPKAICGIIASKNVLTT 60

DB 2 QDWLTFQKKHLTNRVDVDCNNIMSTNLFHCKDKNTFYSPPEPKAICGIIASKNVLTT 61

OY 61 FEFLSDCNVTSRPCKYKLLKSTNFCVTCENQAPVHFVGHC 104

DB 62 SEFLSDCNVTSRPCKYKLLKSTNFCVTCENQAPVHFVGHC 105

RESULT 5

AAV28879

ID AAV28879 standard; Protein; 127 AA.

AC AAV28879;

XX 25-JAN-2000 (first entry)

DT

DE Rana pipiens Clone 5a1b ribonuclease.

XX

KM Rana pipiens ribonuclease Clone 5a1b; RapLr1; covalently bound; RNase;

KM LL2 antibody; ligand binding moiety; CD22; cancerous B cell; onconase;

KM Kaposi's Sarcoma; human chorionic gonadotropin; hCG; cancer;

KW recombinant ribonuclease; frog; signal peptide; cytotoxic fusion protein;

KW autoimmune disease.

XX

OS Rana pipiens.

OS

FT Key Location/Qualifiers

FT Peptide 1..23

FT /label="Signal-peptide"

FT /note="Putative"

FT Protein 24..127

XX

XX /label="Rana_pipiens_Clone_5a1b_ribonuclease"

XX

PN W09950398-A2.

XX

PD 07-OCT-1999.

XX

XX 26-MAR-1999; 99WO-US06641.

PF

XX 27-MAR-1998; 98US-0079751.

PR

XX (USSH) US DEPT HEALTH & HUMAN SERVICES.

PA

PI Newton DL, Rybak SM;

XX

DR WPI: 1999-610847/52.

DR N-PSDB; AAZ08136.

XX

XX New recombinant ribonucleases, used for killing target cells, e.g. for

PT treating cancers, viral infections or autoimmune diseases

XX

PS Disclosure; Page 69; 71pp; English.

XX

CC The present sequence is a Rana pipiens Clone 5a1b ribonuclease (RapLr1).

CC It is encoded by Clone 5a1b cDNA obtained from Rana pipiens liver mRNA

CC library. It exhibits differences with Onconase (RTM) at amino acid

CC residues 11, 20, 85 and 103. Carboxy terminal end of RapL1 has a
 CC covalently bound ligand binding moiety, which can be a LL2 antibody
 CC directed against CD22 on cancerous B cells or human chorionic
 CC gonadotrophin (hCG) effective against Kaposi's Sarcoma cells. Recombinant
 CC ribonucleases can be expressed in bacteria without an N-terminal
 CC methionine due to the presence of a signal peptide that is cleaved by
 CC bacteria. The soluble expression of ribonuclease allows the proteins to
 CC be fused in-frame with ligand binding moieties to form cytotoxic fusion
 CC proteins. They can be used for treatment of cancer and autoimmune
 CC diseases.

SO Sequence 127 AA;

Query Match 98.3%; Score 569; DB 20; Length 127;
 Best Local Similarity 98.1%; Pred. No. 5.4e-62;
 Matches 102; Conservative 1; Mismatches 1; Indels 0; Gaps 0;

OY 1 QDWLTFQKKHLTNTRDVCNNIISTNLFHCKDKNTFTYSRPPVKAICKGIASKNVLT 60
 |||||||
 DB 24 QDWLTFQKKHLTNTRDVCNNIISTNLFHCKDKNTFTYSRPPVKAICKGIASKNVLT 83
 |||||||

OY 61 EFYLSDCNVTSRPCKYKRLKSTNFCVTCENQAPVHFVGVC 104
 |||||||

DB 84 EFYLSDCNVTSRPCKYKRLKSTNFCVTCENQAPVHFVGVC 127
 |||||||

RESULT 6
 AAY28870
 ID AAY28870 standard; Protein: 104 AA.

AC AAY28870;
 XX
 DT 25-JAN-2000 (first entry)

DE Recombinant RapL1 Gln1Ser amino acid sequence.

XX
 KW Recombinant Rana pipiens ribonuclease; RapL1 Gln1Ser; covalently bound;
 KW LL2 antibody; ligand binding moiety; CD22; cancerous B cell; frog;
 KW Kaposi's sarcoma; human chorionic gonadotrophin; hCG; signal peptide;
 KW recombinant ribonuclease; cytotoxic fusion protein; cancer; RNase;
 KW autoimmune disease.

XX
 OS Rana pipiens.
 OS Synthetic.

XX
 FH Key Location/Qualifiers
 FT MISC-difference 1 /note= "Wild type Gln replaced with Ser"
 FT

XX
 PN W09950398-A2.
 XX
 PD 07-OCT-1999.
 XX
 PE 26-MAR-1999; 99WO-US06641.
 XX
 PR 27-MAR-1998; 98US-0079751.
 XX
 PA (USSH) US DEPT HEALTH & HUMAN SERVICES.
 XX
 PI Newton DL, Rybak SM;
 XX
 DR WPI; 1999-610847/52.
 DR N-PSDB; AA208128.
 XX
 PT New recombinant ribonucleases, used for killing target cells, e.g. for
 PT treating cancers, viral infections or autoimmune diseases -
 XX
 PS Claim 34; Page 60; 71pp; English.

XX
 CC The present sequence is a recombinant Rana pipiens ribonuclease (RapL1)
 CC protein with Gln1Ser. Carboxy terminal end of recombinant RapL1 has a
 CC covalently bound ligand binding moiety, which can be a LL2 antibody
 CC directed against CD22 on cancerous B cells or human chorionic

CC gonadotrophin (hCG) effective against Kaposi's sarcoma cells. Recombinant
 CC ribonucleases can be expressed in bacteria without an N-terminal
 CC methionine due to the presence of a signal peptide that is cleaved by
 CC bacteria. The soluble expression of ribonuclease allows the proteins to
 CC be fused in-frame with ligand binding moieties to form cytotoxic fusion
 CC proteins. They can be used for treatment of cancer and autoimmune
 CC diseases.

SO Sequence 104 AA;

Query Match 97.4%; Score 564; DB 20; Length 104;
 Best Local Similarity 98.1%; Pred. No. 1.7e-61;
 Matches 101; Conservative 1; Mismatches 1; Indels 0; Gaps 0;

OY 2 DMLTFQKKHLTNTRDVCNNIISTNLFHCKDKNTFTYSRPPVKAICKGIASKNVLT 61
 |||||||
 DB 2 DMLTFQKKHLTNTRDVCNNIISTNLFHCKDKNTFTYSRPPVKAICKGIASKNVLT 61
 |||||||

OY 62 EFYLSDCNVTSRPCKYKRLKSTNFCVTCENQAPVHFVGVC 104
 |||||||

DB 62 EFYLSDCNVTSRPCKYKRLKSTNFCVTCENQAPVHFVGVC 104
 |||||||

RESULT 7
 AAY28871
 ID AAY28871 standard; Protein: 105 AA.

AC AAY28871;
 XX
 DT 25-JAN-2000 (first entry)

DE Recombinant Met(-1) RapL1 Gln1Ser amino acid sequence.

XX
 KW Recombinant Met(-1) Rana pipiens ribonuclease Gln1Ser; RapL1; CD22;
 KW covalently bound; LL2 antibody; ligand binding moiety; cancerous B cell;
 KW Kaposi's sarcoma; human chorionic gonadotrophin; hCG; signal peptide;
 KW recombinant ribonuclease; cytotoxic fusion protein; cancer; frog;
 KW autoimmune disease; RNase.

XX
 OS Rana pipiens.
 OS Synthetic.

XX
 FH Key Location/Qualifiers
 FT MISC-difference 1 /note= "Met not found in wild type RapL1"
 FT
 FT MISC-difference 2 /note= "Wild type Gln replaced with Ser"
 FT

XX
 PN W09950398-A2.
 XX
 PD 07-OCT-1999.
 XX
 PE 26-MAR-1999; 99WO-US06641.
 XX
 PR 27-MAR-1998; 98US-0079751.
 XX
 PA (USSH) US DEPT HEALTH & HUMAN SERVICES.
 XX
 PI Newton DL, Rybak SM;
 XX
 DR WPI; 1999-610847/52.
 DR N-PSDB; AA208129.
 XX
 PT New recombinant ribonucleases, used for killing target cells, e.g. for
 PT treating cancers, viral infections or autoimmune diseases -
 XX
 PS Claim 34; Page 61; 71pp; English.

XX
 CC The present sequence is a recombinant Rana pipiens ribonuclease (RapL1)
 CC protein with Met at position 1 and Gln1Ser. Carboxy terminal end of
 CC recombinant RapL1 has a covalently bound ligand binding moiety, which
 CC can be a LL2 antibody directed against CD22 on cancerous B cells or human
 CC chorionic gonadotrophin (hCG) effective against Kaposi's sarcoma cells.


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XX      frog; ribonuclease; ranpirinase; RNase.
KW      Rana pipiens.
XX      XX
OS      Rana pipiens.
XX      XX
FH      Key Location/Qualifiers
FT      Modified-site 1
XX      /note= "this Gln is autocyclised to pyroglutamic acid"
XX      US6175003-B1.
XX      16-JAN-2001.
XX      PD
XX      10-SEP-1999; 99US-0394268.
XX      PF
XX      10-SEP-1999; 99US-0394268.
XX      PR
XX      10-SEP-1999; 99US-0394268.
XX      XX
PA      (ALFA-) ALFACELL CORP.
XX      PI
XX      Saxena SK;
XX      DR
XX      WPI; 2001-167808/17.
XX      PT
XX      New nucleic acids encoding a ribonuclease (Rnase), useful for the
XX      precise targeting of Rnase to a predetermined cell receptor .
XX      PS
XX      Claim 1; Columns 5-6; 7pp; English.
XX      CC
XX      The present sequence represents a frog ribonuclease protein (ranpirinase)
XX      (Rnase). The specification describes a synthetic ribonuclease protein,
XX      in which the addition of cysteine in the ribonuclease facilitates the
XX      chemical linking of a targeting molecule by the single reactive
XX      sulfhydryl group. The specification also describes a method for the
XX      production of ranpirinase using DNA technology instead of processing
XX      biological material. The re-engineering of the protein molecule allows
XX      easier attachment to a targeting molecule thereby making it possible for
XX      the ribonuclease to be delivered to a particular cell receptor where it
XX      might be most effective.
XX      CC
XX      Sequence 104 AA;
XX      SQ
XX      Query Match 94.5%; Score 547; DB 22; Length 104;
XX      Best Local Similarity 94.2%; Pred. No. 2,1e-59;
XX      Matches 98; Conservative 3; Mismatches 3; Indels 0; Gaps 0;
XX      QY
XX      1 QDWLTFQKHILTNTRDYDCNNILSTNLFHCKDKNTFTYSRPEPYKAICKGIASKNVLT 60
XX      |||||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX      Db 1 QDWLTFQKHILTNTRDYDCNNILSTNLFHCKDKNTFTYSRPEPYKAICKGIASKNVLT 60
XX      QY 61 FEFYLSDCNVTSRPCRYKLRKSTNTFCVTCENAPVHFVGSHC 104
XX      |||||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX      Db 61 FEFYLSDCNVTSRPCRYKLRKSTNTFCVTCENAPVHFVGSHC 104
XX      RESULT 11
XX      AAW35126
XX      ID AAW35126 standard; Protein: 379 AA.
XX      AC AAW35126;
XX      XX
XX      DT 20-APR-1998 (first entry)
XX      DE R. pipiens recombinant Rnase ronc fusion protein 2.
XX      KW Rnase A; ribonuclease; cytotoxic; onconase; ronc; immunofusion;
XX      tumour cell growth; frog.
XX      OS Rana pipiens.
XX      OS Synthetic.
XX      PN WO9731116-A2.
XX      PD 28-AUG-1997.

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XX      19-FEB-1997; 97WO-US02588.
XX      PF
XX      21-FEB-1996; 96US-0011800.
XX      PR
XX      (USSH ) US DEPT HEALTH & HUMAN SERVICES.
XX      PA
XX      Boque L, Newton DL, Rybak SM, Wlodawer A;
XX      PI
XX      WPI; 1997-435168/40.
XX      DR
XX      N-PSDB; AAT94964.
XX      DR
XX      Ribonuclease molecules based on native Onconase - used for killing
XX      PT cells, particularly tumour cells
XX      PT
XX      Disclosure; Page 68; 90pp; English.
XX      PS
XX      Sequences AAW35125 to AAW35135 represent recombinant fusion proteins
XX      CC (ronc) which are modifications of the Rnase Onconase (RTM) (ronc). Such
XX      CC novel ribonuclease molecules are highly cytotoxic and can be used alone
XX      CC or to form chemical conjugates or to target recombinant immunofusions.
XX      CC They are used particularly for decreasing tumour cell growth. They can
XX      CC also be used for cell separation in vitro by selectively killing unwanted
XX      CC types of cells, e.g. in bone marrow prior to transplantation into a
XX      CC patient undergoing marrow ablation by radiation, or for killing leukaemia
XX      CC cells or T-cells that would cause graft versus host disease. The toxins
XX      CC can also be used to selectively kill unwanted cells in culture. The new
XX      CC ribonucleases have increased cytotoxic activity compared to ronc and
XX      CC also lower immunogenicity in humans.
XX      CC
XX      Sequence 379 AA;
XX      SQ
XX      Query Match 94.5%; Score 547; DB 18; Length 379;
XX      Best Local Similarity 94.2%; Pred. No. 1,1e-58;
XX      Matches 98; Conservative 3; Mismatches 3; Indels 0; Gaps 0;
XX      QY
XX      1 QDWLTFQKHILTNTRDYDCNNILSTNLFHCKDKNTFTYSRPEPYKAICKGIASKNVLT 60
XX      |||||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX      Db 26 QDWLTFQKHILTNTRDYDCNNILSTNLFHCKDKNTFTYSRPEPYKAICKGIASKNVLT 85
XX      QY 61 FEFYLSDCNVTSRPCRYKLRKSTNTFCVTCENAPVHFVGSHC 104
XX      |||||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX      Db 86 FEFYLSDCNVTSRPCRYKLRKSTNTFCVTCENAPVHFVGSHC 129
XX      RESULT 12
XX      AAW30302
XX      ID AAW30302 standard; Protein: 104 AA.
XX      AC AAW30302;
XX      XX
XX      DT 09-JUN-1998 (first entry)
XX      DE Recombinant onc protein.
XX      KW Onc; onconase; ribonuclease; frog; antitumour; pancreatic cancer;
XX      human immunodeficiency virus type-1; HIV1; replication.
XX      OS Rana pipiens.
XX      OS Synthetic.
XX      OS
XX      FH Key Location/Qualifiers
XX      FT Modified-site 1
XX      FT /note= "pyroglutamic acid; especially
XX      FT 2-pyrrolidone-5-carboxylic acid or
XX      FT 5-oxo-2-pyrrolidinecarboxylic acid"
XX      XX
XX      PN WO9738112-A1.
XX      PD 16-OCT-1997.
XX      PF 04-APR-1997; 97WO-US05675.

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CC	electric point of 9.5 - 10.5, a blocked N-terminal gp. and is free
CC	of carbonyl groups. It is active against certain cancer cells. The
CC	combination of the protein and (2-1-p-dimethylaminoethoxyphenyl)-1,
CC	2-diphenyl-1-butene) citrate salt (tamoxifen) is much more bio-
CC	active than the separate entities against human pancreatic ASPC-1
CC	adenocarcinoma, and the combination of protein and (10-[3-(4-methyl
CC	piperazin-1-yl)-propyl]-2-trifluoromethylphenothiazine (stelazine)
CC	is much more reactive than the separate entities against human lung
CC	A-549 carcinoma. Activity has also been shown against human sub-
CC	maxillary epidermoid carcinoma A-253 cells, human ovarian adeno-
CC	carcinoma NIH-OVCAR-3 cells, human leukemic HL-60 cells, human
CC	COCO 320 DM cells, human LOX melanoma and human lung squamous car-
CC	cinoma HT-520 cells.
xx	
SQ	Sequence 104 AA;
Query Match	94.0%; Score 544; DB 12; Length 104;
Best Local Similarity	93.3%; Pred. No. 5e-59;
Matches	97; Conservative 4; Mismatches 3; Indels 0; Gaps 0;
QY	1 ODWLTPCKKHLITNRDVCNNILSTNLFHCKDKMTFTYSREPPKAICGIIASKNVLT 60
Dd	: : : : : : : : : :
1	EDWLTFFKRKHITNTRDDCDIMISTNFLFHKDKMTFTYSRPEPVKAIKGIIASKNVLT 60
QY	61 FEFYLSDCNVTSRPCKRYKLKKSTMYTCVTCENQAPVHPFGVGHC 104
Dd	: : : : : : : :
61	SEFLSDCNVTSRPCRKYLKKSTMKFCVTCENQAPVHPFGVGSC 104
RESULT 14	
AAR47303	
ID	AAR47303 standard; protein: 104 AA.
AC	AAR47303;
DT	09-SEP-1994 (first entry)
XX	
XX	ONCOMASE (pharmaceutical protein).
DE	
XX	Onconase; pharmaceutical; protein: adenocarcinoma; treatment;
KM	cisplatin; melphalan; Adriamycin; ovarian cancer; ovary.
XX	
OS	Synthetic.
XX	
PN	WO9403197-A.
PD	17-FEB-1994.
XX	
Pf	02-JUL-1993; 93WO-USO6357.
PR	30-JUL-1992; 92US-0921180.
XX	
PA	(ALFA-) ALFACEL CORP.
PI	
ARD	Ardelt WJ, Mikulski SM;
DR	WPL: 1994-065396/08.
XX	
PT	Pharmaceutical contg. Cisplatin, Melphalan or Adriamycin - active
PT	In-vitro against OVCAR-3 human ovarian adenocarcinoma cells
PS	
PS	Claim 7; Page 13; 18pp; English.
XX	
CC	This pharmaceutical protein (ONCOMASE) is used in the production of
CC	a bioactive pharmaceutical composition also comprising one of
CC	Cisplatin (cis-diamminedichloroplatinum), Melphalan, (4-[bis-(2-
CC	chloroethyl)amino]-L-phenylamine) or Adriamycin (Doxorubicin HCl).
CC	The composition has bioactivity In vitro against OVCAR-3 human
CC	ovarian adenocarcinoma cells.
XX	
SQ	Sequence 104 AA;
Query Match	94.0%; Score 544; DB 15; Length 104;

